# PATENT COOPERATION TREATY

# **PCT**

### INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference 9792350-0044	FOR FURTHER ACTION	· see Notifi (Form PC below.	cation of Transmittal of International Search Report T/ISA/220) as well as, where applicable, item 5			
International application No. PCT/US02/11586	International filing date (day/ma 12 April 2002 (12.04.2002)	onth/year)	(Earliest) Priority Date (day/month/year) 20 June 2001 (20.06.2001)			
Applicant LEWIS GRUBER						
This international search report h applicant according to Article 18	as been prepared by this International . A copy is being transmitted to the In	Searching Aternational	Authority and is transmitted to the Bureau.			
This international search report c	onsists of a total of $3$ sheets. upanied by a copy of each prior art does	rument cite	d in this report.			
<ol> <li>Basis of the Report</li> <li>With regard to the language, the international search was carried out on the basis of the international application in the language in which it was filed, unless otherwise indicated under this item.</li> </ol>						
b. With regard to any nuc	the international search was carried out on the basis of a translation of the international application furnished to this Authority (Rule 23.1(b)).  b. With regard to any nucleotide and/or amino acid sequence disclosed in the international application, the international search was carried out on the basis of the sequence listing:					
contained in the inter	national application in written form.					
filed together with th	filed together with the international application in computer readable form.					
furnished subsequent	furnished subsequently to this Authority in written form.					
furnished subsequent	furnished subsequently to this Authority in computer readable form.					
the statement that the international applicat	the statement that the subsequently furnished written sequence listing does not go beyond the disclosure in the international application as filed has been furnished.  the statement that the information recorded in computer readable form is identical to the written sequence listing has been furnished.					
the statement that the been furnished.						
2. Certain claims were	Certain claims were found unsearchable (See Box I).					
3. Unity of invention is	Unity of invention is lacking (See Box II).					
4. With regard to the title,						
1 .	s submitted by the applicant.					
the text has been estal	the text has been established by this Authority to read as follows:					
	•		:			
5. With regard to the abstract,						
	s submitted by the applicant.					
may, within one mont Authority.	blished, according to Rule 38.2(b), by the from the date of mailing of this internated in the first thing of the international first the first things are the first things and the first things are th	is Authority utional searc	as it appears in Box III. The applicant the report, submit comments to this			
6. The figure of the drawings to	The figure of the drawings to be published with the abstract is Figure No.					
as suggested by the ap	plicant.	<u> </u>	None of the figures			
because the applicant	failed to suggest a figure.					
because this figure bet	tter characterizes the invention.					

## INTERNATIONAL SEARCH REPORT

International application No.

PCT/US02/11586

<del></del>		101/0502/11500	
IPC(7) US CL	SSIFICATION OF SUBJECT MATTER  : C12Q 1/68; C12M 1/34; C07H 21/02, 21/0 : 435/6, 287.2; 536/22.1, 23.1, 24.3, 24.31, 2  International Patent Classification (IPC) or to both	4.32, 24.33; 359/566	
B. FIEL	DS SEARCHED		***************************************
Minimum do U.S. : 4	ocumentation searched (classification system followers) 35/6, 287.2; 536/22.1, 23.1, 24.3, 24.31, 24.32, 24.32	d by classification symbols) 4.33; 359/566	
Documentati	on searched other than minimum documentation to the	he extent that such documents are included	in the fields searched
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) Please See Continuation Sheet			
	UMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where a		Relevant to claim No.
X	US 5,776,674 A (ULMER) 07 July 1998, (07.07.	1998), see whole document especially	1-87, 155-157
Y	abstract, col. 1 lines 45-62, col. 5 lines 17-30 & col. 7 lines 27-44, see entire document. US 6,055,106 A (GRIER et al) 25 April 2000 (25.04.200), see whole document especially col. 3 lines 60-65, col.4 lines 56-65 & col. 5 lines 53-64, see entire document.		88-154
Burther	doguments are listed in the continues of Pau G		
	documents are listed in the continuation of Box C.	See patent family annex.	
"A" document	defining the general state of the art which is not considered to be ar relevance	Ister document published after the inter- date and not in conflict with the applica principle or theory underlying the inven	tion but cited to understand the tion
"E" earlier app	plication or patent published on or after the international filing date	"X" document of particular relevance; the considered novel or cannot be considered	laimed invention cannot be ed to involve an inventive step
"L" document establish ti specified)	which may throw doubts on priority claim(s) or which is cited to be publication date of another citation or other special reason (as	"Y" document of particular relevance; the ci considered to involve an inventive step combined with one or more other such	when the document is
"O" document	referring to an oral disclosure, use, exhibition or other means	being obvious to a person skilled in the	art
priority da		"&" document member of the same patent fa	mily .
Date of the actual completion of the international search		Date of mailing of the international search	ch report
15 September 2002 (15.09.2002)		<b>01</b> OCT 2002	• .
Com: Box F Wash	iling address of the ISA/US missioner of Patents and Trademarks	Authorized officer  LOCATION ISELL - HON  Telephone No. 703-308-0196	ris for
	D10 (cored to a) (T-1, 1000)		

Form PCT/ISA/210 (second sheet) (July 1998)

•	PCT/US02/11586		
INTERNATIONAL SEARCH REPORT			
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	·		
A 10 10 AB			
Continuation of B. FIELDS SEARCHED Item 3:			
EAST-USAT, USPGPUB, EPO, JPO, DERWENT, IBM-TDB, STN-BIOSIS, MEDLINE, CANCERLIT, BIOTECHDS, LIFESCI, CAPLUS, EM	ID A CE		
search terms: optical trap, tweezer, beamlet, beam multiple,	BASE		
• •			

Form PCT/ISA/210 (second sheet) (July 1998)

## 'ATENT COOPERATION TRF

INTERNATIONAL PRELIMINARY EX	AMINING AUTHORITY	?		
To: JEFFREY F. CRAFT SONNENSCHEIN NATH & ROSENTHAL P.O. BOX 06180 WACKER DRIVE STATION, SEAR TOWER		PCT		
CHICAGO, IL 60606	OWER		WRITTEN OPINION	
			(PCT Rule 66)	
		Date of Mailing (day/month/year) 26 JUN 2003		
Applicant's or agent's file reference		REPLY DUE within 2 months/days from		
097923500045 International application No.	International filing date	(day/month/year)	the above date of mailing Priority date (day/month/year)	
1				
PCT/US02/11586 International Patent Classification (IPC)	or both national classifica	002) tion and IPC	20 June 2001 (20.06.2001)	
	-		: 435/6, 287.2; 536/22.1, 23.1, 24.3, 24.31,	
ARRYX, INC.				
<ol> <li>This written opinion is the <u>fin</u></li> <li>This opinion contains indicate</li> </ol>			liminary Examining Authority.	
	· ·			
II Priority	I Basis of the opinion			
III Non-establishment of opinion with regard to novelty, inventive step and industrial applicability				
IV Lack of unity of invention				
V Reasoned statement under Rule 66.2 (a)(ii) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement				
VI Certain document	s cited	·		
VII Certain defects in	VII Certain defects in the international application			
VIII Certain observation	ons on the international ap	plication		
3. The applicant is hereby invit	ed to reply to this opinion	· Y		
When? See the time i		o applicant may, befo	re the expiration of that time limit, request	
How? By submitting a written reply, accompanied, where appropriate, by amendments, according to Rule 66.3. For the form and the language of the amendments, see Rules 66.8 and 66.9.				
Also For an additional opportunity to submit amendments, see Rule 66.4.  For the examiner's obligation to consider amendments and/or arguments, see Rule 66.4 bis.  For an informal communication with the examiner, see Rule 66.6				
		·	e established on the basis of this opinion.	
4. The final date by which the international preliminary examination report must be established according to Rule 69.2 is: 20 October 2003 (20.10-2003)				
Name and mailing address of the IPEA/US				
Mail Stop PCT, Attn: IPEA/US				
P.O. Box 1450				
Alexandria, Virginia 22313-1450  Facsimile No. (703)305-3230  Telephone No. 703-308-0196				
Form PCT/IPEA/408 (cover sheet)(July 1998)				
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### WRITTEN OPINION

Inter application No.
PCT/US02/11586

I.	Basis	f the opinion	
1.	With r	egard to the elements of the international application:*	
		the international application as originally filed the description: toages 1-17, as originally filed toages NONE, filed with the demand toages NONE, filed with the letter of	
-	I I I I I I I I I I I I I I I I I I I	the claims:  pages 18-35, as originally filed pages NONE, as amended (together with any statement) under Article 19 pages NONE, filed with the demand pages NONE, filed with the letter of	_• ·
		he drawings:  pages 1-5, as originally filed  pages NONE, filed with the demand  pages NONE, filed with the letter of	e e
	I	he sequence listing part of the description:  pages NONE, as originally filed  pages NONE, filed with the demand  pages NONE, filed with the letter of	
	langua These	regard to the language, all the elements marked above were available or furnished to this A ge in which the international application was filed, unless otherwise indicated under this iter elements were available or furnished to this Authority in the following language	mwhich is:
		the language of a translation furnished for the purposes of international search (under Rule 23 the language of publication of the international application (under Rule 48.3(b)). The language of the translation furnished for the purposes of international preliminary examinates of 5.2 and/or 55.3).	
		regard to any nucleotide and/or amino acid sequence disclosed in the international applicate n was drawn on the basis of the sequence listing:	tion, the written
	f   f   f   f	ontained in the international application in printed form.  iled together with the international application in computer readable form.  urnished subsequently to this Authority in written form.  urnished subsequently to this Authority in computer readable form.  The statement that the subsequently furnished written sequence listing does not go beyond the other maternational application as filed has been furnished.	e disclosure in the
	T $\square$	The statement that the information recorded in computer readable form is identical to the writer been furnished.	tten sequence listing
4.	г [] ] ] ]	The amendments have resulted in the cancellation of:  the description, pages NONE the claims, Nos. NONE the drawings, sheets/fig NONE	
5.	☐ ī b	his opinion has been drawn as if (some of) the amendments had not been made, since they have been eyond the disclosure as filed, as indicated in the Supplemental Box (Rule 70.2(c)).	considered to go
	Replace	ment sheets which have been furnished to the receiving Office in response to an invitation under Article n as "originally filed."	? 14 are referred to in

#### WRITTEN OPINION

International application No. PCT/US02/11586

V. Reasoned statement under Rule 66.2(a)(ii) with regard to novelty, inventive step or industrial applicability; citati ns and explanations supporting such statement

1.	STATEMENT			
	Novelty (N)	Claims	88-154	YES
		Claims	1-87,155-157	NO
	Inventive Step (IS)	Claims	NONE	YES
		Claims	1-157	NO
	Industrial Applicability (IA)	Claims	1-157	YES
		Claims	NONE	NO

#### 2. CITATIONS AND EXPLANATIONS

Claims 1-87 and 155-157 lack novelty under PCT Article 33(2) as being anticipated by (US 5,776,674, 07 July 1998). Ulmer teach the apparatus and method of using an movable optical trap to capture bound and unbound multiple probes and trapping the probes and tracking the probe as it interacts with a target (see whole document esp. abstract, col. 1 lines 45-62, col. 5 lines 17-30 and col. 7 lines 27-44). They teach a multi-position scanning laser trap which is used to translate multiple single molecule complexes in parallel as in an array fashion for increased throughput. Particles can be moved independently (see col. 12 lines 37-45). They teach that Multibeam photonic Tweezers for use of multiposition traps (see col. 12 line 49). They teach the use of laser tweezers traps (see col.1 line 49 and col. 1 line 39). A computer is operably connected to apparatus (see col.12 line 21). They teach the use of video camera (see col. 2 line 44). They teach that optical trap may include objective lens and beam (see figure 2A-2H and col. 2 line 50-52). They teach the use of various lasers and fluorescent dyes to detect the spectrum of the label (see col. 10 line 62 -col.11 line 21). They teach the assay of several biological and chemical material such as antigen and antibody, DNA oligomucleotide extensions (see col. 5 lines 17-30 and col.6 line 15-42). They teach the use of subcells or regions in which the particles are moved (see Figure 3A & col. 4 lines 30-40).

Claims 88-154 lack an inventive step under PCT Article 33(3) as being obvious over Ulmer et al (US 5,776,674, 07 July 1998) in view of Grier et al (US 6,055,106 25 April 2000).

Ulmer teach the apparatus and method of using an movable optical trap to capture bound and unbound multiple probes and trapping the probes and tracking the probe as it interacts with a target (see whole document especially abstract, col. 1 lines 45-62, col. 5 lines 17-30 and col. 7 lines 27-44). They teach a multi-position scanning laser trap which is used to translate multiple single molecule complexes in parallel as in an array fashion for increased throughput. Particles can be moved independently (see col. 12 lines 37-45). They teach that Multibeam photonic Tweezers for use of multiposition traps (see col. 12 line 49). They teach the use of laser tweezers traps (see col. 1 2 line 49 and col. 1 line 39). A computer is operably connected to apparatus (see col.12 line 21). They teach the use of video camera (see col.2 line 44). They teach that optical trap may include objective lens and beam (see figure 2A-2H and col. 2 line 50-52). They teach the use of various lasers and fluorescent dyes to detect the spectrum of the label (see col. 10 line 62 - col.11 line 21). They teach the assay of several biological and chemical material such as antigen and antibody, DNA oligonucleotide extensions (see col. 5 lines 17-30 and col.6 line 15-42). They teach the use of subcells or regions in which the particles are moved (see Figure 3A and col. 4 lines 30-40).

Grier et al teach the formation of various beam patterns to create arrays of optical traps (see whole document and col. 3 line 60-65). They teach the use of diffractive optical element which can include computer generated holograms which split the input light beam into patterns of different regions (see col. 4 lines 56-65). They teach the beams then enter the back aperature 24 of objective lens 20 (see Figure 3). They also teach the use of dichroic beamsplitter to split beam (see col. 5 line 53-64). They teach dynamically translating of traps (see col.5 lines 22-44).

One of ordinary skill in the art would have been motivated to apply Grier et al's teaching of multiple optical trap to Ulmer et al's method of assaying in order to increase the throughput of samples.

Form PCT/IPEA/408 (Box V) (July 1998)

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In the stational application No. PCT/US02/11586

Supplemental Box (To be used when the space in any of the preceding boxes is not sufficient)				
TIME LIMIT:  The time limit set for response to a Written Opinion may not be extended. 37 CFR 1.484(d). Any response received after the expiration of the time limit set in the Written Opinion will not be considered in preparing the International Preliminary Examination Report.				
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